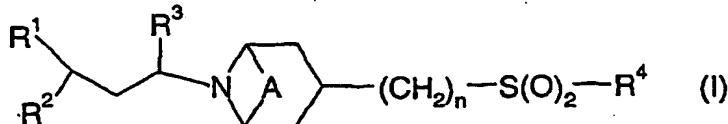


CLAIMS

1. A compound of formula (I):



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wherein:

A is absent or is  $(\text{CH}_2)_2$ ;

$\text{R}^1$  is  $\text{C}(\text{O})\text{NR}^{10}\text{R}^{11}$ ,  $\text{C}(\text{O})_2\text{R}^{12}$ ,  $\text{NR}^{13}\text{C}(\text{O})\text{R}^{14}$ ,  $\text{NR}^{15}\text{C}(\text{O})\text{NR}^{16}\text{R}^{17}$ ,  $\text{NR}^{18}\text{C}(\text{O})_2\text{R}^{19}$ , heterocyclyl, aryl or heteroaryl;

$\text{R}^{10}$ ,  $\text{R}^{13}$ ,  $\text{R}^{15}$ ,  $\text{R}^{16}$  and  $\text{R}^{18}$  are hydrogen or  $\text{C}_{1-6}$  alkyl;

10  $\text{R}^{11}$ ,  $\text{R}^{12}$ ,  $\text{R}^{14}$ ,  $\text{R}^{17}$  and  $\text{R}^{19}$  are  $\text{C}_{1-8}$  alkyl (optionally substituted by halo, hydroxy,  $\text{C}_{1-6}$  alkoxy,  $\text{C}_{1-6}$  haloalkoxy,  $\text{C}_{3-6}$  cycloalkyl (optionally substituted by halo),  $\text{C}_{5-6}$  cycloalkenyl,  $\text{S}(\text{C}_{1-4}$  alkyl),  $\text{S}(\text{O})(\text{C}_{1-4}$  alkyl),  $\text{S}(\text{O})_2(\text{C}_{1-4}$  alkyl), heteroaryl, aryl,

15 heteroaryloxy or aryloxy), aryl, heteroaryl,  $\text{C}_{3-7}$  cycloalkyl (optionally substituted by halo or  $\text{C}_{1-4}$  alkyl),  $\text{C}_{4-7}$  cycloalkyl fused to a phenyl ring,  $\text{C}_{5-7}$  cycloalkenyl, or, heterocyclyl (itself optionally substituted by oxo,  $\text{C}(\text{O})(\text{C}_{1-6}$  alkyl),  $\text{S}(\text{O})_k(\text{C}_{1-6}$  alkyl), halo or  $\text{C}_{1-4}$  alkyl); or  $\text{R}^{11}$ ,  $\text{R}^{12}$ ,  $\text{R}^{14}$  and  $\text{R}^{17}$  can also be hydrogen;

20 or  $\text{R}^{10}$  and  $\text{R}^{11}$ , and/or  $\text{R}^{16}$  and  $\text{R}^{17}$  may join to form a 4-, 5- or 6-membered ring which optionally includes a nitrogen, oxygen or sulphur atom, said ring being optionally substituted by  $\text{C}_{1-6}$  alkyl,  $\text{S}(\text{O})_l(\text{C}_{1-6}$  alkyl) or  $\text{C}(\text{O})(\text{C}_{1-6}$  alkyl);

$\text{R}^2$  is phenyl, heteroaryl or  $\text{C}_{3-7}$  cycloalkyl;

$\text{R}^3$  is H or  $\text{C}_{1-4}$  alkyl;

$\text{R}^4$  is heterocyclyl;

n is 1, 2 or 3;

25 aryl, phenyl and heteroaryl moieties are independently optionally substituted by one or more of halo, cyano, nitro, hydroxy,  $\text{OC}(\text{O})\text{NR}^{20}\text{R}^{21}$ ,  $\text{NR}^{22}\text{R}^{23}$ ,  $\text{NR}^{24}\text{C}(\text{O})\text{R}^{25}$ ,  $\text{NR}^{26}\text{C}(\text{O})\text{NR}^{27}\text{R}^{28}$ ,  $\text{S}(\text{O})_2\text{NR}^{29}\text{R}^{30}$ ,  $\text{NR}^{31}\text{S}(\text{O})_2\text{R}^{32}$ ,  $\text{C}(\text{O})\text{NR}^{33}\text{R}^{34}$ ,  $\text{CO}_2\text{R}^{36}$ ,  $\text{NR}^{37}\text{CO}_2\text{R}^{38}$ ,  $\text{S}(\text{O})_q\text{R}^{39}$ ,  $\text{OS}(\text{O})_2\text{R}^{49}$ ,  $\text{C}_{1-6}$  alkyl (optionally mono-substituted by  $\text{S}(\text{O})_2\text{R}^{50}$  or  $\text{C}(\text{O})\text{NR}^{51}\text{R}^{52}$ ),  $\text{C}_{2-6}$  alkenyl,  $\text{C}_{2-6}$  alkynyl,  $\text{C}_{3-10}$  cycloalkyl,  $\text{C}_{1-6}$  haloalkyl,  $\text{C}_{1-6}$  alkoxy( $\text{C}_{1-6}$ )alkyl,  $\text{C}_{1-6}$  alkoxy,  $\text{C}_{1-6}$  haloalkoxy, phenyl, phenyl( $\text{C}_{1-4}$ )alkyl, phenoxy, phenylthio, phenylS(O), phenylS(O)<sub>2</sub>, phenyl( $\text{C}_{1-4}$ )alkoxy, heteroaryl,

heteroaryl(C<sub>1-4</sub>)alkyl, heteroaryloxy or heteroaryl(C<sub>1-4</sub>)alkoxy; wherein any of the immediately foregoing phenyl and heteroaryl moieties are optionally substituted with halo, hydroxy, nitro, S(C<sub>1-4</sub> alkyl), S(O)(C<sub>1-4</sub> alkyl), S(O)<sub>2</sub>(C<sub>1-4</sub> alkyl), S(O)<sub>2</sub>NH<sub>2</sub>, S(O)<sub>2</sub>NH(C<sub>1-4</sub> alkyl), S(O)<sub>2</sub>N(C<sub>1-4</sub> alkyl)<sub>2</sub>, cyano, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkoxy, C(O)NH<sub>2</sub>,

5 C(O)NH(C<sub>1-4</sub> alkyl), C(O)N(C<sub>1-4</sub> alkyl)<sub>2</sub>, CO<sub>2</sub>H, CO<sub>2</sub>(C<sub>1-4</sub> alkyl), NHC(O)(C<sub>1-4</sub> alkyl), NHS(O)<sub>2</sub>(C<sub>1-4</sub> alkyl), CF<sub>3</sub> or OCF<sub>3</sub>;

unless otherwise stated heterocyclyl is optionally substituted by C<sub>1-6</sub> alkyl [optionally substituted by phenyl {which itself optionally substituted by halo, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkoxy, cyano, nitro, CF<sub>3</sub>, OCF<sub>3</sub>, (C<sub>1-4</sub> alkyl)C(O)NH, S(O)<sub>2</sub>NH<sub>2</sub>, C<sub>1-4</sub> alkylthio, S(O)(C<sub>1-4</sub> alkyl) or S(O)<sub>2</sub>(C<sub>1-4</sub> alkyl)} or heteroaryl {which itself optionally substituted by halo, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkoxy, cyano, nitro, CF<sub>3</sub>, (C<sub>1-4</sub> alkyl)C(O)NH, S(O)<sub>2</sub>NH<sub>2</sub>, C<sub>1-4</sub> alkylthio, S(O)(C<sub>1-4</sub> alkyl) or S(O)<sub>2</sub>(C<sub>1-4</sub> alkyl)} or heteroaryl {which itself optionally substituted by halo, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkoxy, cyano, nitro, CF<sub>3</sub>, OCF<sub>3</sub>, (C<sub>1-4</sub> alkyl)C(O)NH, S(O)<sub>2</sub>NH<sub>2</sub>, C<sub>1-4</sub> alkylthio, S(O)(C<sub>1-4</sub> alkyl) or S(O)<sub>2</sub>(C<sub>1-4</sub> alkyl)}], phenyl {optionally substituted by halo, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkoxy, cyano, nitro, CF<sub>3</sub>, (C<sub>1-4</sub> alkyl)C(O)NH, S(O)<sub>2</sub>NH<sub>2</sub>, C<sub>1-4</sub> alkylthio, S(O)(C<sub>1-4</sub> alkyl) or S(O)<sub>2</sub>(C<sub>1-4</sub> alkyl)}, heteroaryl {optionally substituted by halo, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkoxy, cyano, nitro, CF<sub>3</sub>, (C<sub>1-4</sub> alkyl)C(O)NH, S(O)<sub>2</sub>NH<sub>2</sub>, C<sub>1-4</sub> alkylthio, S(O)(C<sub>1-4</sub> alkyl) or S(O)<sub>2</sub>(C<sub>1-4</sub> alkyl)}, S(O)<sub>2</sub>NR<sup>40</sup>R<sup>41</sup>, C(O)R<sup>42</sup>, C(O)<sub>2</sub>(C<sub>1-6</sub> alkyl) (such as tert-butoxycarbonyl), C(O)<sub>2</sub>(phenyl(C<sub>1-2</sub> alkyl)) (such as benzyloxycarbonyl), C(O)NHR<sup>43</sup>, S(O)<sub>2</sub>R<sup>44</sup>, NHS(O)<sub>2</sub>NHR<sup>45</sup>, NHC(O)R<sup>46</sup>, NHC(O)NHR<sup>47</sup> or NHS(O)<sub>2</sub>R<sup>48</sup>, provided none of these last four substituents is linked to a ring nitrogen;

k, l and q are, independently, 0, 1 or 2;

R<sup>20</sup>, R<sup>22</sup>, R<sup>24</sup>, R<sup>26</sup>, R<sup>27</sup>, R<sup>29</sup>, R<sup>31</sup>, R<sup>33</sup>, R<sup>37</sup>, R<sup>40</sup> and R<sup>51</sup> are, independently, hydrogen or C<sub>1-6</sub> alkyl;

25 R<sup>21</sup>, R<sup>23</sup>, R<sup>25</sup>, R<sup>28</sup>, R<sup>30</sup>, R<sup>32</sup>, R<sup>34</sup>, R<sup>36</sup>, R<sup>38</sup>, R<sup>39</sup>, R<sup>41</sup>, R<sup>42</sup>, R<sup>43</sup>, R<sup>44</sup>, R<sup>45</sup>, R<sup>46</sup>, R<sup>47</sup>, R<sup>48</sup>,

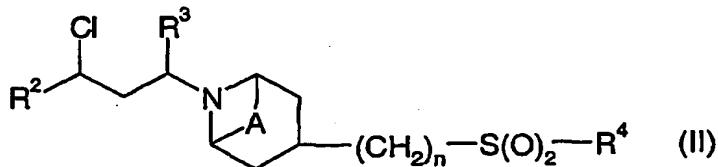
R<sup>49</sup>, R<sup>50</sup> and R<sup>52</sup> are, independently, C<sub>1-6</sub> alkyl (optionally substituted by halo, hydroxy, C<sub>1-6</sub> alkoxy, C<sub>1-6</sub> haloalkoxy, C<sub>3-6</sub> cycloalkyl, C<sub>5-6</sub> cycloalkenyl, S(C<sub>1-4</sub> alkyl), S(O)(C<sub>1-4</sub> alkyl), S(O)<sub>2</sub>(C<sub>1-4</sub> alkyl), heteroaryl, phenyl, heteroaryloxy or phenoxy), C<sub>3-7</sub> cycloalkyl, phenyl or heteroaryl; wherein any of the immediately foregoing phenyl and heteroaryl moieties are optionally substituted with halo, hydroxy, nitro, S(C<sub>1-4</sub> alkyl),

30 S(O)(C<sub>1-4</sub> alkyl), S(O)<sub>2</sub>(C<sub>1-4</sub> alkyl), S(O)<sub>2</sub>NH<sub>2</sub>, S(O)<sub>2</sub>NH(C<sub>1-4</sub> alkyl), S(O)N(C<sub>1-4</sub> alkyl)<sub>2</sub>, cyano, C<sub>1-4</sub> alkyl, C<sub>1-4</sub> alkoxy, C(O)NH<sub>2</sub>, C(O)NH(C<sub>1-4</sub> alkyl), C(O)N(C<sub>1-4</sub> alkyl)<sub>2</sub>, CO<sub>2</sub>H, CO<sub>2</sub>(C<sub>1-4</sub> alkyl), NHC(O)(C<sub>1-4</sub> alkyl), NHS(O)<sub>2</sub>(C<sub>1-4</sub> alkyl), C(O)(C<sub>1-4</sub> alkyl), CF<sub>3</sub> or OCF<sub>3</sub>;

$R^{21}$ ,  $R^{23}$ ,  $R^{25}$ ,  $R^{28}$ ,  $R^{30}$ ,  $R^{34}$ ,  $R^{35}$ ,  $R^{36}$ ,  $R^{41}$ ,  $R^{42}$ ,  $R^{43}$ ,  $R^{45}$ ,  $R^{46}$ ,  $R^{47}$  and  $R^{52}$  may additionally be hydrogen;  
or a pharmaceutically acceptable salt thereof or a solvate thereof.

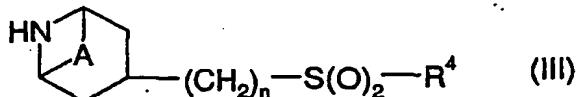
- 5    2. A compound as claimed in claim 1 wherein  $R^1$  is  $NR^{13}C(O)R^{14}$ , wherein  $R^{13}$  and  $R^{14}$  are as defined in claim 1.
- 10    3. A compound as claimed in claim 1 or 2 wherein  $R^1$  is optionally substituted aryl or optionally substituted heteroaryl, wherein the optional substituents are as recited in claim 1.
- 15    4. A compound as claimed in claim 1, 2 or 3 wherein  $R^1$  is optionally substituted heterocyclyl.
- 20    5. A compound as claimed in any one of the preceding claims wherein  $R^2$  is phenyl optionally substituted by halo or  $CF_3$ .
- 25    6. A compound as claimed in any one of the preceding claims wherein  $R^3$  is hydrogen.
7. A compound as claimed in any one of the preceding claims wherein  $R^4$  is heterocyclyl optionally substituted by oxo, halogen, cyano, hydroxy,  $C_{1-6}$  alkyl (itself optionally substituted by halogen, hydroxy, cyano or  $C_{1-4}$  alkoxy),  $C_{2-4}$  alkenyl,  $CO_2(C_{1-4}$  alkyl),  $S(O)_2(C_{1-4}$  alkyl),  $CH(O)$ ,  $S(O)_2(C_{1-4}$  haloalkyl),  $C(O)(C_{1-4}$  alkyl),  $C(O)(C_{3-6}$  cycloalkyl),  $N(C_{1-4}$  alkyl)<sub>2</sub>,  $C(O)NH_2$ ,  $C(O)N(C_{1-4}$  alkyl)<sub>2</sub> or  $NHC(O)(C_{1-4}$  alkyl).
- 30    8. A compound as claimed in any one of the preceding claims wherein heterocyclyl is piperidinyl, homopiperazinyl, thiomorpholinyl, pyrrolidinyl, piperazinyl, 1,2,3,6-tetrahydropyridinyl, morpholinyl, 2,5-dihydropyrrolyl, azetidinyl, 1,4-oxepanyl, 3-azabicyclo[3.2.1]octan-3-yl, 8-azaspiro[4.5]decanyl or 3-azabicyclo[3.1.0]hex-3-yl.
9. A compound as claimed in any one of the preceding claims wherein A is absent.
10. A compound as claimed in any one of the preceding claims wherein n is 2.

11. A process for preparing a compound as claimed in claim 1, the process comprising:
- when R<sup>1</sup> is an N-linked optionally substituted heterocycle, reacting a compound of formula (II):

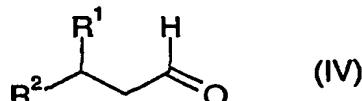


wherein R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, n, A and X are as defined in claim 1, with a compound R<sup>1</sup>H (wherein the H is on a heterocycle ring nitrogen atom) wherein R<sup>1</sup> is as defined in claim 1, in the presence of a suitable base and in a suitable solvent;

- when R<sup>3</sup> is hydrogen, coupling a compound of formula (III):



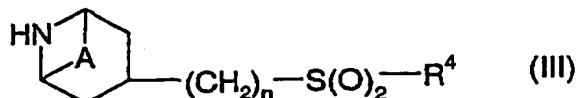
wherein R<sup>4</sup>, n, A and X are as defined in claim 1, with a compound of formula (IV):



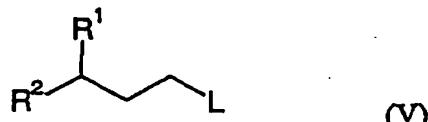
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wherein R<sup>1</sup> and R<sup>2</sup> are as defined in claim 1, in the presence of NaBH(OAc)<sub>3</sub> (wherein Ac is C(O)CH<sub>3</sub>) in a suitable solvent at room temperature; or,

- when R<sup>3</sup> is hydrogen, coupling a compound of formula (III):



wherein R<sup>4</sup>, n, A and X are as defined in claim 1, with a compound of formula (V):



wherein R<sup>1</sup> and R<sup>2</sup> are as defined in claim 1 and L is an activated leaving group, in the presence of a base, in a suitable solvent at a temperature from 60°C up to the boiling point of the solvent.

12. A pharmaceutical composition which comprises a compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof, and a pharmaceutically acceptable adjuvant, diluent or carrier.

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13. A compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof, for use as a medicament.

14. A compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof, in the manufacture of a medicament for use in therapy.

- 10  
15. A method of treating a CCR5 mediated disease state comprising administering to a patient in need of such treatment an effective amount of a compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof.

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